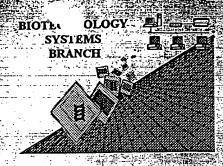
# RAW SEQUENCE LISTING ERROR REPORT



The Biotechnology Systems Branch of the Scientific and Technical Information— Center (STIC) detected errors when processing the following computer readable form:

Application Serial Number: 09/461,580

Source: 1645

Date Processed by STIC: 7-26-00

THE ATTACHED PRINTOUT EXPLAINS DETECTED ERRORS.

PLEASE FORWARD THIS INFORMATION TO THE APPLICANT BY EITHER:

- 1) INCLUDING A COPY OF THIS PRINTOUT IN YOUR NEXT COMMUNICATION TO THE APPLICANT, WITH A NOTICE TO COMPLY or,
- 2) TELEPHONING APPLICANT AND FAXING A COPY OF THIS PRINTOUT, WITH A NOTICE TO COMPLY

FOR FURTHER INFORMATION, PLEASE TELEPHONE MARK SPENCER, 703-308-4212.

TO REDUCE ERRORED SEQUENCE LISTINGS, PLEASE USE THE <u>CHECKER</u> <u>VERSION 3.0 PROGRAM</u>, ACCESSIBLE THROUGH THE U.S. PATENT AND TRADEMARK OFFICE WEBSITE. SEE BELOW:

#### Checker Version 3.0

The Checker Version 3.0 application is a state-of the-art Windows based software program employing a logical and intuitive user-interface to check whether a sequence listing is in compliance with format and content rules. Checker Version 3.0 works for sequence listings generated for the original version of 37 CFR §§1.821 – 1.825 effective October 1, 1990 (old rules) and the revised version (new rules) effective July 1, 1998 as well as World Intellectual Property Organization (WIPO) Standard ST.25.

Checker Version 3.0 replaces the previous DOS-based version of Checker, and is Y2K-compliant. Checker allows public users to check sequence listings in Computer Readable form (CRF) before submitting them to the United States Patent and Trademark Office (USPTO). Use of Checker prior to filing the sequence listing is expected to result in fewer errored sequence listings, thus saving time and money.

Checker Version 3.0 can be down loaded from the USPTO website at the following address: http://www.uspto.gov/web/offices/pac/checker

## Ray Sequence Listing Error Sux nary

### ERROR DETECTED SUGGESTED CORRECTION SERIAL NUMBER:

ATTN: NEW RULES CASES: PLEASE DISREGARD ENGLISH "ALPHA" HEADERS, WHICH WERE INSERTED BY PTO SOFTWARE The number/text at the end of each line "wrapped" down to the next line. Wrapped Nucleics This may occur if your file was retrieved in a word processor after creating it. Please adjust your right margin to .3, as this will prevent "wrapping". The amino acid number/text at the end of each line "wrapped" down to the next line. Wrapped Aminos This may occur if your file was retrieved in a word processor after creating it. Please adjust your right margin to .3, as this will prevent "wrapping". J Incorrect Line Length The rules require that a line not exceed 72 characters in length. This includes spaces. Misaligned Amino Acid The numbering under each 5th amino acid is misaligned. This may be caused by the use of tabs Numbering between the numbering. It is recommended to delete any tabs and use spacing between the numbers. Non-ASCII This file was not saved in ASCII (DOS) text, as required by the Sequence Rules. Please ensure your subsequent submission is saved in ASCII text so that it can be processed. Sequence(s) \_\_\_\_ contain n's or Xaa's which represented more than one residue. Variable Length As per the rules, each n or Xaa can only represent a single residue. Please present the maximum number of each residue having variable length and indicate in the (ix) feature section that some may be missing. Patentin ver. 2.0 "bug" A "bug" in PatentIn version 2.0 has caused the <220>-<223> section to be missing from amino acid . Normally, Patentin would automatically generate this section from the previously coded nucleic acid sequence. Please manually copy the relevant <220>-<223> section to the subsequent amino acid sequence. This applies primarily to the mandatory <220>-<223> sections for Artificial or Unknown sequences. missing. If intentional, please use the following format for each skipped sequence: **Skipped Sequences** Sequence(s) (OLD RULES) (2) INFORMATION FOR SEQ ID NO:X: (i) SEQUENCE CHARACTERISTICS:(Do not insert any headings under "SEQUENCE CHARACTERISTICS") (xi) SEQUENCE DESCRIPTION:SEQ ID NO:X: This sequence is intentionally skipped Please also adjust the "(iii) NUMBER OF SEQUENCES:" response to include the skipped sequence(s). Skipped Sequences Sequence(s) missing. If intentional, please use the following format for each skipped sequence. (NEW RULES) <210> sequence id number <400> sequence id number Use of n's or Xaa's Use of n's and/or Xaa's have been detected in the Sequence Listing. (NEW RULES) Use of <220> to <223> is MANDATORY if n's or Xaa's are present. In <220> to <223> section, please explain location of n or Xaa, and which residue n or Xaa represents. \_\_ Use of <213>Organism are missing this mandatory field or its response. (NEW RULES) Sequence(s)\_\_ are missing the <220>Feature and associated headings. Use of <220> to <223> is MANDATORY if <213>ORGANISM is "Ártificial" or "Unknown" Please explain source of genetic material in <220> to <223> section. (See "Federal Register," 6/01/98, Vol. 63, No. 104, pp. 29631-32) (Sec. 1.823 of new Rules) Please do not use "Copy to Disk" function of PatentIn version 2.0. This causes a corrupted Patentin ver. 2.0 "bug" file, resulting in missing mandatory numeric identifiers and responses (as indicated on raw sequence listing).

Instead, please use "File Manager" or any other means to copy file to floppy disk.



### **Does Not Comply** Corrected Diskette Needec

1645

RECEIVED

RAW SEQUENCE LISTING

PATENT APPLICATION: US/09/461,580

DATE: 07/26/2000

TIME: 14:00:15

AUG 16 2000

Input Set : A:\00501618000.txt

Output Set: N:\CRF3\07262000\I461580.raw

TECH CENTER 1600/2900

See P. 5, 7

```
4 <110> APPLICANT: Guarente, Leonard
          Imai, Shin-ichiro
          Armstrong, Christopher
 8 <120> TITLE OF INVENTION: METHODS FOR IDENTIFYING AGENTS WHICH
         ALTER HISTINE PROTEIN ACETYLATION, DECREASE AGING, INCREASE
         LIFESPAN
12 <130> FILE REFERENCE: 0050.1618-000
14 <140> CURRENT APPLICATION NUMBER: 09/461,580
15 <141> CURRENT FILING DATE: 1999-12-15
17 <160> NUMBER OF SEQ ID NOS: 35
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29 Ala Ala Ala Ala Met Glu Ala Ala Ser Gln Pro Ala Asp Glu Pro Leu
30 25 30
31 Arg Lys Arg Pro Arg Arg Asp Gly Pro Gly Leu Gly Arg Ser Pro Gly 32 \phantom{-}35\phantom{+}40\phantom{+}45\phantom{+}
33 Glu Pro Ser Ala Ala Val Ala Pro Ala Ala Ala Gly Cys Glu Ala Ala
34 50 55 60
35 Ser Ala Ala Ala Pro Ala Ala Leu Trp Arg Glu Ala Ala Gly Ala Ala
36 65 70 75 80
37 Ala Ser Ala Glu Arg Glu Ala Pro Ala Thr Ala Val Ala Gly Asp Gly 38 \phantom{\bigg|}85\phantom{\bigg|}90\phantom{\bigg|}95\phantom{\bigg|}
                     85
39 Asp Asn Gly Ser Gly Leu Arg Arg Glu Pro Arg Ala Ala Asp Asp Phe
40 100 105 110
41 Asp Asp Asp Glu Glu Glu Glu Asp Glu Ala Ala Ala Ala Ala Ala Ala 42 115 120 125
43 Ala Ala Ala Ile Gly Tyr Arg Asp Asn Leu Leu Leu Thr Asp Gly Leu
44 130 135 140
45 Leu Thr Asn Gly Phe His Ser Cys Glu Ser Asp Asp Asp Asp Arg Thr
46 145 150 160
47 Ser His Ala Ser Ser Ser Asp Trp Thr Pro Arg Pro Arg Ile Gly Pro 48 165 170 175
49 Tyr Thr Phe Val Gln Gln His Leu Met Ile Gly Thr Asp Pro Arg Thr 50 180 185 190
51 Ile Leu Lys Asp Leu Leu Pro Glu Thr Ile Pro Pro Pro Glu Leu Asp
          195
                                  200
                                                           205
53 Asp Met Thr Leu Trp Gln Ile Val Ile Asn Ile Leu Ser Glu Pro Pro 54 210 215 220
55 Lys Arg Lys Lys Arg Lys Asp Ile Asn Thr Ile Glu Asp Ala Val Lys 56 225 230 235 240
57 Leu Leu Gln Glu Cys Lys Lys Ile Ile Val Leu Thr Gly Ala Gly Val
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DATE: 07/26/2000 TIME: 14:00:15 RAW SEQUENCE LISTING PATENT APPLICATION: US/09/461,580

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Output Set: N:\CRF3\07262000\1461580.raw

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	Ser	Val	Ser		Gly	Ile	Pro	Asp		Arg	Ser	Arg	Asp	270	iie	Tyr
60		_	_	260			D1	D	265	<b>.</b>	D	2	D		212	Wat
62	Ala	Arg	Leu 275	Ala	var	Asp	Pne	280	ASP	Leu	PIO	Asp	285	GIH	Ald	met
63	Phe	Asp	Ile	Glu	Tyr	Phe	Arg	Lys	Asp	Pro	Arg	Pro	Phe	Phe	Lys	Phe
64		290			_		295					300				
65	Ala	Lys	Glu	Ile	Tyr	Pro	Gly	Gln	Phe	Gln	Pro	Ser	Leu	Cys	His	Lys
	305	-				310					315					320
67	Phe	Ile	Ala	Leu	Ser	Asp	Lys	Glu	Gly	Lys	Leu	Leu	Arg	Asn	Tyr	Thr
68					325					330					335	
69	Gln	Asn	Ile	Asp	Thr	Leu	Glu	Gln	Val	Ala	Gly	Ile	Gln	Arg	Ile	Leu
70				340					345					350		
71	Gln	Cys	His	Gly	Ser	Phe	Ala	$\mathtt{Thr}$	Ala	Ser	Cys	Leu	Ile	Cys	Lys	Tyr
72			355					360					365			
73	Lys	Val	Asp	Cys	Glu	Ala	Val	Arg	Gly	Asp	Ile	Phe	Asn	Gln	Val	Val
74		370					375					380				
75	Pro	Arg	Cys	Pro	Arg	Cys	Pro	Ala	Asp	Glu	Pro	Leu	Ala	Ile	Met	
	385					390					395					400
77	Pro	Glu	Ile	Val	Phe	Phe	Gly	Glu	Asn	Leu	Pro	Glu	Gln	Phe		Arg
78					405					410					415	
79	Ala	Met	Lys	Tyr	Asp	Lys	Asp	Glu		Asp	Leu	Leu	Ile	Val	Ile	Gly
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82			435					440				_	445	_		_
	His		Val	Pro	Gln	Ile		He	Asn	Arg	Glu	Pro	Leu	Pro	His	Leu
84		450					455			_	_	460			_	~ 1
		Phe	Asp	Val	Glu		Leu	GIĀ	Asp	Cys		Val	ire	ite	Asn	
	465	_		_	_	470		~ 1			475	<b>-</b>	<b></b>	G	3	480
	Leu	Cys	HIS	Arg		GIY	GTĀ	GIU	туг		гĀг	Leu	Cys	Cys	495	PIO
88	** - 1			<b>a</b>	485	+1-	m1	<b>61</b>	T	490	Dwo	7	Dwo	C1 n		Clu
	val	гля	Leu	500	GIU	ire	TIII	GIU	505	PIO	PIO	Arg	PIO	510	цуз	GIU
90	T	17.0.1	114 -		Com	C1	T 011	Dro		mb~	Dro	Leu	uic		Cor	Clu
92	Leu	val	515	Leu	ser	Giu	пец	520	FIO	1111	FIU	neu	525	110	561	O.Lu
	N c n	cor		cor	Dro	C111	λra		Val	Dro	Cln	Asp		Ser	Va 1	Tle
94	vaħ	530	361	JCI	110	Giu	535	1111	vu1	110	01	540	001	501		
	Δla		Leu	Va1	Asn	Gln		Thr	Asn	Asn	Asn	Val	Asn	Asp	Leu	Glu
	545	1111	пси	vui	nop	550	2114	****	11511		555					560
		Ser	Glu	Ser	Ser		Val	Glu	Glu	Lvs		Gln	Glu	Val	Gln	Thr
98		501	014	001	565	0,75				570					575	
	Ser	Ara	Asn	Val		Asn	Tle	Asn	Val	-	Asn	Pro	Asp	Phe	Lvs	Ala
100		5		580					585					590		
		l G1s	, Sei			. Ala	a Asr	o Lvs			ı Ar	Thi	s Se	r Va.	L Ala	ı Glu
10:			59					600				•	60			
		r Va			CVS	Tr	Pro			ı Lei	ı Ala	a Lys	Gl	ı Glı	ı Ile	e Ser
104		610		, -1.	-1.		615					620				
				ı Glı	ı Glv	Ası			. Lei	ı Phe	e Vai	l Pro	Pro	) Ası	a Arc	y Tyr
	625			-		630					63					640
								•								

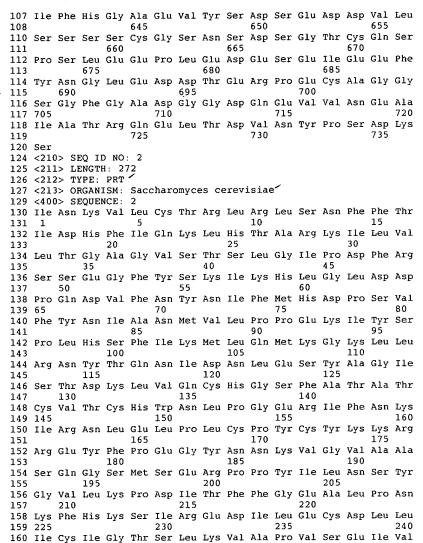




RAW SEQUENCE LISTING DATE: 07/26/2000 PATENT APPLICATION: US/09/461,580 TIME: 14:00:15

Input Set : A:\00501618000.txt

Output Set: N:\CRF3\07262000\I461580.raw









DATE: 07/26/2000 TIME: 14:00:15 RAW SEQUENCE LISTING PATENT APPLICATION: US/09/461,580

Input Set : A:\00501618000.txt
Output Set: N:\CRF3\07262000\I461580.raw

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	Asn	Met	Val	Pro		His	Val	Pro	Gln		Leu	Ile	Asn	Arg		Pro
. 163				260					265					270		
	<210	)> SI	it os		. 3											
		l> Li	-													
		2> T														
						hard	าพงะเ	-s ce	erev	isiae	_ /					
		)> S!				,,,,,,,,,,	J.II. <sub>2</sub> O.									
			_			Ser	Thr	Ara	Leu	Ara	Leu	Pro	Asn	Phe	Asn	Thr
173	1		-,-		5					10					15	
		Asp	His	Phe	Thr	Ala	Thr	Leu	Arg	Asn	Ala	Lvs	Lvs	Ile	Leu	Val
175				20					25				•	30		
176	Leu	Thr	Glv	Ala	Glv	Val	Ser	Thr	Ser	Leu	Gly	Ile	Pro	Asp	Phe	Arq
177			35		2			40			•		45	•		•
178	Ser	Ser	Glu	Glv	Phe	Tvr	Ser	Lvs	Ile	Arq	His	Leu	Gly	Leu	Glu	Asp
179		50				•	55	•		-		60	•			•
180	Pro	Gln	Asp	Val	Phe	Asn	Leu	Asp	Ile	Phe	Leu	Gln	Asp	Pro	Ser	Val
181			•			70		-			75					80
182	Phe	Tyr	Asn	Ile	Ala	His	Met	Val	Leu	Pro	Pro	Glu	Asn	Met	Tyr	Ser
183					85					90					95	
184	Pro	Leu	His	Ser	Phe	Ile	Lys	Met	Leu	Gln	Asp	Lys	Gly	Lys	Leu	Leu
185				100					105					110		
186	Arg	Asn	Tyr	Thr	Gln	Asn	Ile	Asp	Asn	Leu	Glu	Ser	Tyr	Ala	Gly	Ile
187			115					120					125			
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189		130					135					140				
	-	Val	Thr	Cys	His	-	Gln	Ile	Pro	Gly		Lys	Ile	Phe	Glu	
	145					150					155					160
	Ile	Arg	Asn	Leu		Leu	Pro	Leu	Cys		Tyr	Cys	Tyr	Gln	_	Arg
193	_		_		165		_	_		170	_				175	_
	Lys	Gln	Tyr		Pro	Met	Ser	Asn	Gly	Asn	Asn	Thr	Val		Thr	Asn
195			-1	180	•		-1-		185	<b>a</b>	m	G1	1	190	T	D
	тте	Asn		Asn	Ser	Pro	TTE		Lys	ser	Tyr	GTÅ		ren	ьұs	Pro
197		14-4	195	nh-	nh.	C1	<b>~1</b>	200	T	D==0	Com	7	205	TT	T	mla sa
198	Asp		Thr	Pne	Pne	GŢĀ	215	ATG	Leu	PIO	ser	220	rne	нтз	Lys	THE
	т1.	210	T ***	A a n	T10	T ou		Cuc	Asp	T 011	T OIL		Cuc	T10	C111	Thr
	225	Arg	гÃ2	ASP	TTE	230	GIU	Cys	ASP	цец	235	116	Cys	116	GIY	240
		LOU	Luc	17 - 1	λla		Wa 1	Car	Glu	T10		Acn	Mot	V = 1	Dro	
203	361	пеп	цуз	Val	245	110	Val	Jei	GIU	250	VUL	ASII	ricc	var	255	DCI
	Hic	Va 1	Dro	Gln		T.011	Tle	Δen	Arg		Met				233	
205		• 42		260	110	LÇU			265	p						
	05															
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DATE: 07/26/2000 RAW SEQUENCE LISTING PATENT APPLICATION: US/09/461,580 TIME: 14:00:15

Input Set : A:\00501618000.txt

Output Set: N:\CRF3\07262000\I461580.raw

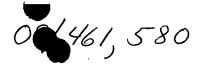
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218 Ile Ile Val Leu, Thr Gly Ala Gly Val Ser Val Ser Cys Gly Ile Pro
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221 Asp Phe Arg Ser Arg Asp Gly Ile Tyr Ala Arg Leu Ala Val Asp Phe
       50
                            55
223 Pro Asp Leu Pro Asp Pro Gln Ala Met Phe Asp Ile Glu Tyr Phe Arg
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225 Lys Asp Pro Arg Pro Phe Phe Lys Phe Ala Lys Glu Ile Tyr Pro Gly
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                                        90
227 Gln Phe Gln Pro Ser Leu Cys His Lys Phe Ile Ala Leu Ser Asp Lys
             100
                                   105
                                                        110
229 Glu Gly Lys Leu Leu Arg Asn Tyr Thr Gln Asn Ile Asp Thr Leu Glu
230 115 120 125
231 Gln Val Ala Gly Ile Gln Arg Ile Leu Gln Cys His Gly Ser Phe Ala
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232
     130
                                               140
233 Thr Ala Ser Cys Leu Ile Cys Lys Tyr Lys Val Asp Cys Glu Ala Val
234 145 150 155 160
235 Arg Gly Asp Ile Phe Asn Gln Val Val Pro Arg Cys Pro Arg Cys Pro
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                   165
237 Ala Asp Glu Pro Leu Ala Ile Met Lys Pro Glu Ile Val Phe Phe Gly
238
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                                                        190
239 Glu Asn Leu Pro Glu Gln Phe His Arg Ala Met Lys Tyr Asp Lys Asp
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                                200
241 Glu Val Asp Leu Leu Ile Val Ile Gly Ser Ser Leu Lys Val Arg Pro
                                                220
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245 Ile Asn Arg Glu Pro
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252 <213> ORGANISM: Escherichia coli
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                                40
261 Pro Gly Leu Val Gln Thr Phe Tyr Asn Ala Arg Arg Gln Gln Leu Gln
       50
                           55
263 Gln Pro Glu Ile Gln Pro Asn Ala Ala His Leu Ala Leu Ala Asn Leu
                        70
                                            75
265 Lys Lys Arg Leu Ala Ile Ala Phe Leu Leu Val Thr Gln Asn Ile Asp
                   85
                                        90
267 Asn Leu His Glu Arg Ala Gly Asn Arg Asn Ile Ile Gln Met His Gly
```

F. Y. 1.

Please Note:

Use of n and/or Xaa have been detected in the Sequence Listing. Please review the Sequence Listing to ensure that a corresponding explanation is presented in the <220> to <223> fields of each sequence which presents at least one n or Xaa.

p. 7



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	0> 3!								•							_				
	1> 2					Y	, ,		a 4	6	-1.6		(2)	20	> <	22	3	>		
	2> PI													_						
<21.	3> u:	nknov	wn		7	Fec	Li	ر <i>و</i> ر	t	, ()	EX	ola.	'n	orig	sin	o f	1	Unk	nou	179
< 400	)> 3!	5									,							~ <i>Ca</i> =		•-
Thr 1	_	Pro	Arg	Thr 5	Arg	Gly	His	Gly	Thr 10	Glu	Gln	Ser	Val	Thr 15	Val		۷	orga	7131	-,
Pro	Arg	Ala	Pro 20	Thr	Pro	Phe	Pro	Trp 25	Pro	Ser	Arg	Thr	Asp 30	Ser	Asp		S	୧୧	+	£ 12
Ser	Asp	Thr 35	Glu	Gly	Gly	Ala	Thr 40	Gly	Gly	Glu	Ala	Glu 45	Met	Asp	Phe	$\sim$		E	610	<i>/</i>
Leu	Arg 50	Asn	Leu	Phe	Thr	Gln 55	Thr	Leu	Gly	Leu	Gly 60	Ser	Gln	Lys	Glu		•	ee E	<u>۸</u>	,
Arg 65	Leu	Leu	Asp	Glu	Leu 70	Thr	Leu	Glu	Gly	Val 75	Thr	Arg	Tyr	Met	Gln 80		> C	smm Sh	7	, L
Ser	Glu	Arg	Cys	Arg 85	Lys	Val	Ile	Cys	Leu 90	Val	Gly	Ala	Gly	Ile 95	Ser			Sh	66	/
Thr	Ser	Ala	Gly 100	Ile	Pro	Asp	Phe	Arg 105	Ser	Pro	Ser	Thr	Gly 110	Leu	Tyr					
Ala	Asn	Leu 115	Glu	Lys	Tyr	His	Leu 120	Pro	Tyr	Pro	Glu	Ala 125	Ile	Phe	Glu					
Ile	Ser 130	Tyr	Phe	Lys	Lys	His 135	Pro	Glu	Pro	Phe	Phe 140	Ala	Leu	Ala	Lys					
Glu 145	Leu	Tyr	Pro	G1y	Gln 150	Phe	Lys	Pro	Thr	Ile 155	Cys	His	Tyr	Phe	Ile 160					
Arg	Leu	Leu	Lys	Glu 165	Lys	Gly	Leu	Leu	Leu 170	Arg	Cys	Tyr	Thr	G1n 175	Asn					
Ile	Asp	Thr	Leu 180	Glu	Arg	Val	Ala	Gly 185	Leu	Glu	Pro	Gln	Asp 190	Leu	Val					
Glu	Ala	His 195	Gly	Thr	Phe	Tyr	Thr 200	Ser	His	Cys	Val	Asn 205	Thr	Ser	Cys					
Arg	Lys 210	Glu	Tyr	Thr	Met	Gly 215	Trp	Met	Lys	Glu	Lys 220	Ile	Ser	Gln	Lys					
Gln 225	Leu	Pro	Gly	Val	Ser 230	Ser	Val													





DATE: 07/26/2000 TIME: 14:00:16

VERIFICATION SUMMARY
PATENT APPLICATION: US/09/461,580

Input Set : A:\00501618000.txt
Output Set: N:\CRF3\07262000\1461580.raw

L:1110 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:27
L:1261 M:361 W: Invalid Split Codon, Sequence data for SEQ ID#: 34
L:1270 M:258 W: Mandatory Feature missing, <220> FEATURE:
L:1270 M:258 W: Mandatory Feature missing, <223> OTHER INFORMATION: